## SUPPLEMENTAL MATERIAL

## Competitive Inhibition of Thyroidal Uptake of Dietary Iodide by Perchlorate Does Not Describe Perturbations in Rat Serum Total T<sub>4</sub> and TSH

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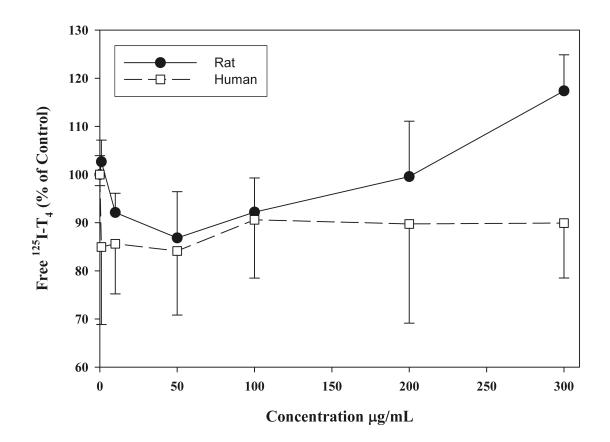
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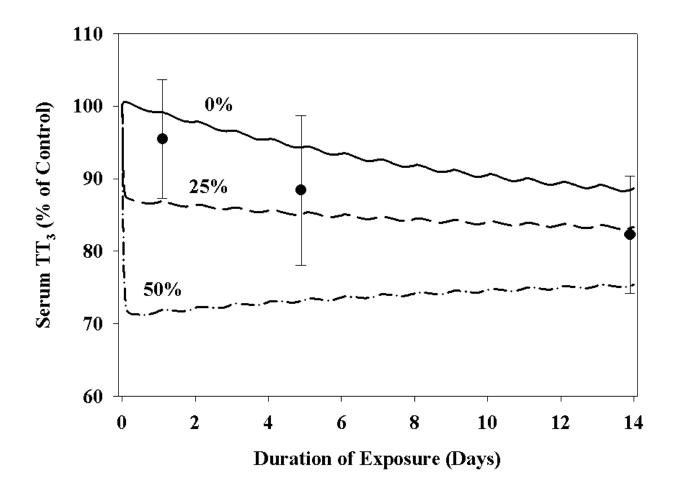
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Supplemental Material, Figure 1. Effects of perchlorate on free  $^{125}$ I-T<sub>4</sub> in serum as determined by *in-vitro* equilibrium dialysis. Results are expressed as a percent of control +SE (rat serum) and – SE (human serum). N = 6 for the control measurement and N = 4 for the perchlorate concentrations.



Supplemental Material, Figure 2. Model predictions of serum  $TT_3$  compared with data following exposure to 10 mg/kg perchlorate per day in drinking water. Model predictions are shown testing 0, 25, or 50% inhibition of thyroid hormone production. Data  $\pm$  SD for 10 mg/kg-day from Yu et al. (2002).



Supplemental Material, Table 1. Model predicted serum T<sub>4</sub> and TSH concentrations as percent of control for a 300g adult rat compared to available literature data for 0-10 mg ClO<sub>4</sub><sup>-</sup>/kg-day (Yu et al. 2002) and 15 mg ClO<sub>4</sub><sup>-</sup>/kg-day (Männistö et al. 1979). This table corresponds to plots shown in Figure 5 of the manuscript, but is reported because of the difficulty in depicting failed model simulations at the low doses.

	1 Day of Perchlorate Exposure					14 Days of Perchlorate Exposure			
Dose of Perchlorate	Serum T <sub>4</sub> (% Control)		Serum	Serum TSH		Serum T <sub>4</sub>		Serum TSH	
(mg/kg-day)			(% Control)		(% Control)		(% Control)		
	Predicted	Observed	Predicted	Observed	Predicted	Observed	Predicted	Observed	
0	100	$100\pm10$	100	$100 \pm 9$	100	$100 \pm 12$	100	$100 \pm 8$	
0.1	100	$94 \pm 11$	100	$162 \pm 15$	100	$97 \pm 10$	100	$124\pm12$	
1	99.9	$89 \pm 8$	100.1	$168\pm17$	99.8	$96 \pm 7$	100.2	$182\pm19$	
3	99.8	$88 \pm 9$	100.2	$196 \pm 19$	99.0	$86 \pm 6$	101.0	$245\pm25$	
10	98.0	$76 \pm 7$	102.1	$273\pm21$	82.8	$84 \pm 9$	120.8	$277\pm30$	
15	94.2ª	$78.3 \pm 6^{a}$	106.1 <sup>a</sup>	$118 \pm 22^{a}$	73.4	Not reported	136.1	$196 \pm 13$	

<sup>&</sup>lt;sup>a</sup> Data (Männistö et al. 1979) and model simulation following 2 days of exposure to 15 mg/kg-day.

## REFERENCES

- Männistö PT, Ranta T, Leppäluoto J. 1979. Effects of methylmercaptoimidazole (MMI), propylthiouracil (PTU), potassium perchlorate (KClO4) and potassium iodide (KI) on the serum concentrations of thyrotrophin (TSH) and thyroid hormones in the rat. Acta Endocrinol 91:271-281.
- Yu KO, Narayanan L, Mattie DR, Godfrey RJ, Todd PN, Sterner TR, et al. 2002. The pharmacokinetics of perchlorate and its effect on the hypothalamus-pituitary-thyroid axis in the male rat. Toxicol Appl Pharmacol 182:148-159.